

US EPA ARCHIVE DOCUMENT



UNITED STATES ENVIRONMENTAL PROTECTION AGENCY
WASHINGTON, D.C. 20460

OFFICE OF
PESTICIDES AND TOXIC SUBSTANCES

MEMORANDUM JAN 27 1983

TO: Richard Mountfort (23)
Registration Division (TS-767)

THRU: Orville E. Paynter, Ph.D.
Chief, Toxicology Branch
Hazard Evaluation Division (TS-769)

SUBJECT: Preliminary Report on Repeat Chronic Rat Study of
Metolachlor CASWELL#188DD

Registrant: Ciba-Geigy Corporation
P.O. Box 11422
Greensboro, North Carolina 27409

Background Information:

In my memo of October 7, 1981, I reviewed a rat chronic feeding study conducted at IBT (IBT No. 622-07926) that was classified as Supplementary Data and which indicated an increase in primary liver tumors in females. The study has recently been repeated at another laboratory, using a similar protocol and dose levels, and the findings regarding liver neoplasia were noted in the letter of December 9, 1982 from Ciba-Geigy to R. Mountfort, Registration Division, EPA. The preliminary report of that study is discussed below.

Discussion/Recommendation:

The preliminary results of the repeat study confirm and strengthen the results of the earlier study. In the second study

an increased incidence of liver neoplasia is observed in both the male and female high dose groups. The tumor incidences are shown in the following table*:

	Sex	Feeding Level (PPM)			
		0	30	300	3000
Proliferative Foci (Neoplastic Nodule)	M	0	2	0	6
	F	0	1	1	5
Hepatocellular (Carcinoma)	M	2	0	2	3
	F	0	0	1	2
Total Animals with Hepatocellular Carcinoma or Neo- plastic Nodules	M	2	2	2	9
	F	0	1	2	7
Total Animals Examined	M	60	57	60	60
	F	59	60	59	60

*This table was presented in the letter of December 9, 1983 from Ciba-Geigy.

It is noted that the tumor incidences for females are similar in the both the original and repeat studies while the rate for high dose males is somewhat greater. Only data on liver neoplasia was available in this preliminary report. Ciba-Geigy has conducted and submitted a risk assessment based on this data; however, a risk assessment based on the repeat study will be performed by EPA after submission and review of the final report for the repeat study.

Chronic Oral, Rats. Conducted at Raltech Scientific Services.

Core Classification: Supplementary Data. (Interim reports on this study were reviewed in my memos of October 7, 1981 and January 26, 1982.) The preliminary findings indicate an increased incidence of liver neoplasia in treated animals.

Gary J. Burin *JDC*
 Gary J. Burin, Toxicologist
 Toxicology Branch
 Hazard Evaluation Division (TS-769) *1/25/83*
1/25/83

REGISTRATION DIVISION DATA REVIEW RECORD
Confidential Business Information - Does Not Contain National Security Information (E.O. 12065)

5151
12/10/82

1. CHEMICAL NAME

metolachlor - "Dual 8E"

2. IDENTIFYING NUMBER

100-597

3. ACTION CODE

575

4. ACCESSION NUMBER

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TO BE COMPLETED BY PM

5. RECORD NUMBER

85121

6. REFERENCE NUMBER

32

7. DATE RECEIVED (EPA)

12/9/82

8. STATUTORY DUE DATE

9. PRODUCT MANAGER (PM)

Mountfort

10. PM TEAM NUMBER

23

14. CHECK IF APPLICABLE

☐ Public Health/Quarantine

☐ Minor Use

☐ Substitute Chemical

☐ Part of IPM

☒ Seasonal Concern

☐ Review Requires Less Than 4 Hours

TO BE COMPLETED BY PCB

11. DATE SENT TO HED/TSS

12/10/82

12. PRIORITY NUMBER

20

13. PROJECTED RETURN DATE

2/10/83

15. INSTRUCTIONS TO REVIEWER

A. HED ☒ Total Assessment - 3(c)(5)

☐ Incremental Risk Assessment - 3(c)(7) and/or E.L. Johnson memo of May 12, 1977.

B. SPRD (Send Copy of Form to SPRD PM)

☐ Chemical Undergoing Active RPAR Review

☐ Chemical Undergoing Active Registration Standards Review

C. ☐ BFSD

D. ☐ TSS/RD

E. ☐ Other

F. INSTRUCTIONS

Preliminary Report of 2nd chronic rat study "liver lesions appeared to result from administration." Ciba-Geigy has included a revised risk estimate. Attention: Gary Burin; Review 1/29/83

16. RELATED ACTIONS

For Petition 2F2720 on 12/2/82 Record # 84710 we requested a new risk assessment

17. 3(c)(1)(D)

☒ Use Any or All Available Information ☐ Use Only Attached Data
Use Only the Attached Data for Formulation and Any or All
☐ Available Information on the Technical or Manufacturing Chemical.

18. REVIEWS SENT TO

☒ TB

☐ EEB

☐ EF

☐ PL

☐ RCB

☐ EFB

☐ CH

☐ BFSD

19. To	TYPE OF REVIEW	NUMBER OF ACTIONS							
		Registration	Petition	EUP	SLN	Sec. 18	Inert	MNR. USE	Other
HED	<input checked="" type="checkbox"/> TOXICOLOGY	1							Returned 2/23/83
	ECOLOGICAL EFFECTS								
	RESIDUE CHEMISTRY								
	ENVIRONMENTAL DATA								
RD/TSS	CHEMISTRY								
	EFFICACY								
	PRECAUTIONARY LABELING								
BFSD	ECONOMIC ANALYSIS								

20. ☐ Label Submitted with Application Attached

21. ☐ Confidential Statement of Formula

22. ☐ Representative Labels Showing Accepted Uses Attached

23. Date Returned to RD (to be completed by HED) 1-27-83

24. Include an Original and 4 (four) Copies of This Completed Form for Each Branch Checked for Review.